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<110> Van Mellaert, Herman Botterman, Johan Van Rie, Jeroen Joos, Henk		
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290 295 300

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agc Ser	ccg Pro	cat His	cta Leu	ctt Leu 325	gat Asp	ttt Phe	cta Leu	gaa Glu	caa Gln 330	ctt Leu	aca Thr	att Ile	ttt Phe	agc Ser 335	gct Ala	1008
tca Ser	tca Ser	cga Arg	tgg Trp 340	agt Ser	aat Asn	act Thr	agg Arg	cat His 345	atg Met	act Thr	tat Tyr	tgg Trp	cgg Arg 350	ggg Gly	cac His	1056
acg Thr	att Ile	caa Gln 355	tct Ser	cgg Arg	cca Pro	ata Ile	gga Gly 360	ggc Gly	gga Gly	tta Leu	aat Asn	acc Thr 365	tca Ser	acg Thr	cat His	1104
ggg Gly	gct Ala 370	acc Thr	aat Asn	act Thr	tct Ser	att Ile 375	aat Asn	cct Pro	gta Val	aca Thr	tta Leu 380	cgg Arg	ttc Phe	gca Ala	tct Ser	1152
cga Arg 385	gac Asp	gtt Val	tat Tyr	agg Arg	act Thr 390	gaa Glu	tca Ser	tat Tyr	gca Ala	gga Gly 395	gtg Val	ctt Leu	cta Leu	tgg Trp	gga Gly 400	1200
att Ile	tac Tyr	ctt Leu	gaa Glu	cct Pro 405	att Ile	cat His	ggt Gly	gtc Val	cct Pro 410	act Thr	gtt Val	agg Arg	ttt Phe	aat Asn 415	ttt Phe	1248
acg Thr	aac Asn	cct Pro	cag Gln 420	aat Asn	att Ile	tct Ser	gat Asp	aga Arg 425	ggt Gly	acc Thr	gct Ala	aac Asn	tat Tyr 430	agt Ser	caa Gln	1296
cct Pro	tat Tyr	gag Glu 435	Ser	cct Pro	gly aaa	ctt Leu	caa Gln 440	tta Leu	aaa Lys	gat Asp	tca Ser	gaa Glu 445	act Thr	gaa Glu	tta Leu	1344
cca Pro	cca Pro 450	gaa Glu	aca Thr	aca Thr	gaa Glu	cga Arg 455	cca Pro	aat Asn	tat Tyr	gaa Glu	tct Ser 460	tac Tyr	agt Ser	cac His	agg Arg	1392
tta Leu 465	Ser	cat His	ata Ile	ggt Gly	ata Ile 470	att Ile	tta Leu	caa Gln	tcc Ser	agg Arg 475	gtg Val	aat Asn	gta Val	ccg Pro	gta Val 480	1440
tat Tyr	tct Ser	tgg Trp	acg Thr	cat His	Arg	agt Ser	gca Ala	gat Asp	cgt Arg 490	Thr	aat Asn	acg Thr	att Ile	gga Gly 495	cca Pro	1488
aat Asn	aga . Arg	ato Ile	acc Thr	Glr	atc Ile	cca Pro	atg Met	gta Val 505	Lys	gca Ala	tcc Ser	gaa Glu	ctt Leu 510	Pro	caa Gln	1536

			gtt Val													1	584
			aat Asn													1	632
gga Gly 545	cca Pro	tta Leu	aca Thr	caa Gln	aga Arg 550	tat Tyr	cgt Arg	ata Ile	gga Gly	ttc Phe 555	cgc Arg	tat Tyr	gct Ala	tca Ser	act Thr 560	1	680
			gat Asp													1	.728
			cta Leu 580													1	.776
aat Asn	ttt Phe	gtg Val 595	aga Arg	cgt Arg	gct Ala	ttt Phe	act Thr 600	aca Thr	cct Pro	ttt Phe	act Thr	ttt Phe 605	aca Thr	caa Gln	att Ile	1	824
caa Gln	gat Asp 610	ata Ile	att Ile	cga Arg	acg Thr	tct Ser 615	att Ile	caa Gln	ggc Gly	ctt Leu	agt Ser 620	gga Gly	aat Asn	ggg Gly	gaa Glu	1	.872
			gat Asp													1	920
gca Ala	gaa Glu	tat Tyr	gat Asp	tta Leu 645	gaa Glu	aga Arg	gcg Ala	caa Gln	gag Glu 650	gcg Ala	gtg Val	aat Asn	gct Ala	ctg Leu 655	ttt Phe	1	.968
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			aag Lys													2	2112
ctc Leu 705	Ser	gat Asp	gaa Glu	aga Arg	aac Asn 710	tta Leu	ctc Leu	caa Gln	gat Asp	cca Pro 715	aac Asn	ttc Phe	aca Thr	tcc Ser	atc Ile 720	2	2160
aat	aag	caa	cca	gac	ttc	ata	tct	act	aat	gag	caa	tcg	aat	ttc	aca	2	2208

Asn	Lys	Gln	Pro	Asp 725	Phe	Ile	Ser	Thr	Asn 730	Glu	Gln	Ser	Asn	Phe 735	Thr	
			_								gga Gly					2256
											aat Asn					2304
											tta Leu 780					2352
											caa Gln					2400
											att Ile					2448
											tcc Ser					2496
tca	att	gaa	agc	cca	atc	aaa	200	+~~	~~~	~ ~ ~	cca	2 2 ±	cas	tac	aca	2544
											Pro					2344
Ser cca	Val cat	Glu 835 ttt	Ser gaa	Pro tgg	Ile aat	Gly	Arg 840 gat	Cys cta	Gly gat	Glu tgt		Asn 845 tgc	Arg aga	Cys gat	Ala gga	2592
Ser cca Pro	Val cat His 850	Glu 835 ttt Phe	Ser gaa Glu gcg	Pro tgg Trp	Ile aat Asn	cct Pro 855	Arg 840 gat Asp	Cys cta Leu cat	Gly gat Asp	Glu tgt Cys	Pro tcc Ser	Asn 845 tgc Cys	Arg aga Arg	Cys gat Asp	Ala gga Gly gtt	
cca Pro gaa Glu 865	cat His 850 aaa Lys	Glu 835 ttt Phe tgt Cys	gaa Glu gcg Ala	tgg Trp cat His	aat Asn cat His 870	cct Pro 855 tcc Ser	Arg 840 gat Asp cat His	Cys cta Leu cat His	gat Asp ttc Phe	tgt Cys tct Ser 875	tcc ser 860	Asn 845 tgc Cys gat Asp	aga Arg att Ile	gat Asp gat Asp	Ala gga Gly gtt Val 880 aag	2592
cca Pro gaa Glu 865 gga Gly	cat His 850 aaa Lys tgc Cys	Glu 835 ttt Phe tgt Cys aca Thr	gaa Glu gcg Ala gac Asp	tgg Trp cat His ttg Leu 885	aat Asn cat His 870 cat His	CCt Pro 855 tcc Ser gag Glu	Arg 840 gat Asp cat His aat Asn	Cys cta Leu cat His cta Leu	gat Asp ttc Phe ggc Gly 890 cta	tgt Cys tct Ser 875 gtg Val	tcc Ser 860 ttg Leu	Asn 845 tgc Cys gat Asp gtg Val	aga Arg att Ile gta Val	gat Asp gat Asp ttc Phe 895	Ala gga Gly gtt Val 880 aag Lys	2592 2640
cca Pro gaa Glu 865 gga Gly att Ile	cat His 850 aaa Lys tgc Cys aag Lys	Glu 835 ttt Phe tgt Cys aca Thr	gaa Glu gcg Ala gac Asp cag Gln 900 cca	tgg Trp cat His ttg Leu 885 gaa Glu	aat Asn cat His 870 cat His	CCt Pro 855 tcc Ser gag Glu cat His	Arg 840 gat Asp cat His aat Asn gca Ala	Cys cta Leu cat His cta Leu aga Arg 905 gca	gat Asp ttc Phe ggc Gly 890 cta Leu	tgt Cys tct ser 875 gtg Val ggg Gly	tcc ser 860 ttg Leu tgg Trp	Asn 845 tgc Cys gat Asp gtg Val ctg Leu	aga Arg att Ile gta Val gaa Glu 910	gat Asp gat Asp ttc Phe 895 ttt Phe	Ala gga Gly gtt Val 880 aag Lys att Ile	2592 2640 2688

930 935 940

cga gta tat aca gag gca aaa gaa gct gtg gat gct tta ttc gta gat Arg Val Tyr Thr Glu Ala Lys Glu Ala Val Asp Ala Leu Phe Val Asp 945 950 955 960	2880
tct caa tat gat aga tta caa gcg gat aca aac atc ggc atg att cat Ser Gln Tyr Asp Arg Leu Gln Ala Asp Thr Asn Ile Gly Met Ile His 965 970 975	2928
gcg gca gat aaa ctt gtt cat cga att cga gag gcg tat ctt tca gaa Ala Ala Asp Lys Leu Val His Arg Ile Arg Glu Ala Tyr Leu Ser Glu 980 985 990	2976
tta cct gtt atc cca ggt gta aat gcg gaa att ttt gaa gaa tta gaa Leu Pro Val Ile Pro Gly Val Asn Ala Glu Ile Phe Glu Glu Leu Glu 995 1000 1005	3024
ggt cac att atc act gca atc tcc tta tac gat gcg aga aat gtc Gly His Ile Ile Thr Ala Ile Ser Leu Tyr Asp Ala Arg Asn Val 1010 1015 1020	3069
gtt aaa aat ggt gat ttt aat aat gga tta aca tgt tgg aat gta Val Lys Asn Gly Asp Phe Asn Asn Gly Leu Thr Cys Trp Asn Val 1025 1030 1035	3114
aaa ggg cat gta gat gta caa cag agc cat cat cgt tct gac ctt Lys Gly His Val Asp Val Gln Gln Ser His His Arg Ser Asp Leu 1040 1045 1050	3159
gtt atc cca gaa tgg gaa gca gaa gtg tca caa gca gtt cgc gtc Val Ile Pro Glu Trp Glu Ala Glu Val Ser Gln Ala Val Arg Val 1055 1060 1065	3204
tgt ccg ggg tgt ggc tat atc ctt cgt gtc aca gcg tac aaa gag Cys Pro Gly Cys Gly Tyr Ile Leu Arg Val Thr Ala Tyr Lys Glu 1070 1075 1080	3249
gga tat gga gag ggc tgc gta acg atc cat gaa atc gag aac aat Gly Tyr Gly Glu Gly Cys Val Thr Ile His Glu Ile Glu Asn Asn 1085 1090 1095	3294
aca gac gaa cta aaa ttt aaa aac cgt gaa gaa gag gaa gtg tat Thr Asp Glu Leu Lys Phe Lys Asn Arg Glu Glu Glu Glu Val Tyr 1100 1105 1110	3339
cca acg gat aca gga acg tgt aat gat tat act gca cac caa ggt Pro Thr Asp Thr Gly Thr Cys Asn Asp Tyr Thr Ala His Gln Gly 1115 1120 1125	3384
aca gct gga tgc gca gat gca tgt aat tcc cgt aat gct gga tat Thr Ala Gly Cys Ala Asp Ala Cys Asn Ser Arg Asn Ala Gly Tyr 1130 1135 1140	3429

gag Glu	gat Asp 1149	Al	a tat a Tyr	ga Gl	a gt u Va	t gat l Asp 1150	Thr	aca Thr	gca Ala	tct Ser	gtt Val 1155	aat Asn	tac Tyr	aaa Lys	3474
ccg Pro	act Thr 1160	Ту	t gaa r Glu	a ga 1 Gl	a gaa u Gli	a acg ı Thr 1165	Tyr	aca Thr	gat Asp	gta Val	aga Arg 1170	aga Arg	gat Asp	aat Asn	3519
cat His	tgt Cys 1175	Gli	a tat ı Tyr	ga As	c aga p Arg	g ggg g Gly 1180	Tyr	gtc Val	aat Asn	tat Tyr	cca Pro 1185	cca Pro	gta Val	cca Pro	3564
gct Ala	ggt Gly 1190	Туз	gtg Val	Th:	a aaa r Lys	gaa Glu 1195	Leu	gaa Glu	tac Tyr	ttc Phe	cca Pro 1200	gaa Glu	aca Thr	gat Asp	3609
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		Va]				ctc Leu 1225				tag					3687
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Asn	His	Ser	Ala 20	Gln	Met	Asp Le	eu Le 25		o As <sub>l</sub>	o Ala	a Arg	Ile 30	Glu	Asp	
Ser		Cys 35	Ile .	Ala	Glu	Gly As 40		n Il	e Asp	p Pro	Phe 45	Val	Ser	Ala	
Ser	Thr 50	Val	Gln '	Thr		Ile As 55	n Il	e Al	a Gly	/ Arg	J Ile	Leu	Gly	Val	
Leu 65	Gly	Val	Pro :	Phe	Ala 70	Gly Gl	n Le	u Al	a Sei 75	? Phe	e Tyr	Ser	Phe	Leu 80	

Val Gly Glu Leu Trp Pro Arg Gly Arg Asp Gln Trp Glu Ile Phe Leu

85 90 95

Glu	His	Val	Glu 100	Gln	Leu	Ile	Asn	Gln 105	Gln	Ile	Thr	Glu	Asn 110	Ala	Arg
Asn	Thr	Ala 115	Leu	Ala	Arg	Leu	Gln 120	Gly	Leu	Gly	Asp	Ser 125	Phe	Arg	Ala
Tyr	Gln 130	Gln	Ser	Leu	Glu	Asp 135	Trp	Leu	Glu	Asn	Arg 140	Asp	Asp	Ala	Arg
Thr 145	Arg	Ser	Val	Leu	His 150	Thr	Gln	Tyr	Ile	Ala 155	Leu	Glu	Leu	Asp	Phe 160
Leu	Asn	Ala	Met	Pro 165	Leu	Phe	Ala	Ile	Arg 170	Asn	Gln	Glu	Val	Pro 175	Leu
Leu	Met	Val	Tyr 180	Ala	Gln	Ala	Ala	Asn 185	Leu	His	Leu	Leu	Leu 190	Leu	Arg
Asp	Ala	Ser 195	Leu	Phe	Gly	Ser	Glu 200	Phe	Gly	Leu	Thr	Ser 205	Gln	Glu	Ile
Gln	Arg 210	Tyr	Tyr	Glu	Arg	Gln 215	Val	Glu	Arg	Thr	Arg 220	Asp	Tyr	Ser	Asp
Tyr 225	Cys	Val	Glu	Trp	Tyr 230	Asn	Thr	Gly	Leu	Asn 235	Ser	Leu	Arg	Gly	Thr 240
Asn	Ala	Ala	Ser	Trp 245	Val	Arg	Tyr	Asn	Gln 250	Phe	Arg	Arg	Asp	Leu 255	Thr
Leu	Gly	Val	Leu 260	Asp	Leu	Val	Ala	Leu 265	Phe	Pro	Ser	Tyr	Asp 270	Thr	Arg
Thr	Tyr	Pro 275	Ile	Asn	Thr	Ser	Ala 280	Gln	Leu	Thr	Arg	Glu 285	Val	Tyr	Thr
Asp	Ala 290	Ile	Gly	Ala	Thr	Gly 295	Val	Asn	Met	Ala	Ser 300	Met	Asn	Trp	Tyr

Asn Asn Asn Ala Pro Ser Phe Ser Ala Ile Glu Ala Ala Ala Ile Arg 305 310 315 Ser Pro His Leu Leu Asp Phe Leu Glu Gln Leu Thr Ile Phe Ser Ala 325 330 335 Ser Ser Arg Trp Ser Asn Thr Arg His Met Thr Tyr Trp Arg Gly His 340 345 Thr Ile Gln Ser Arg Pro Ile Gly Gly Leu Asn Thr Ser Thr His 355 360 Gly Ala Thr Asn Thr Ser Ile Asn Pro Val Thr Leu Arg Phe Ala Ser 370 375 380 Arg Asp Val Tyr Arg Thr Glu Ser Tyr Ala Gly Val Leu Leu Trp Gly 385 390 395 400 Ile Tyr Leu Glu Pro Ile His Gly Val Pro Thr Val Arg Phe Asn Phe 405 410 415 Thr Asn Pro Gln Asn Ile Ser Asp Arg Gly Thr Ala Asn Tyr Ser Gln 420 425 430 Pro Tyr Glu Ser Pro Gly Leu Gln Leu Lys Asp Ser Glu Thr Glu Leu 435 440 445 Pro Pro Glu Thr Thr Glu Arg Pro Asn Tyr Glu Ser Tyr Ser His Arg 455 460 Leu Ser His Ile Gly Ile Ile Leu Gln Ser Arg Val Asn Val Pro Val 465 470 475 Tyr Ser Trp Thr His Arg Ser Ala Asp Arg Thr Asn Thr Ile Gly Pro 485 490 495 Asn Arg Ile Thr Gln Ile Pro Met Val Lys Ala Ser Glu Leu Pro Gln 500 505

Gly Thr Thr Val Val Arg Gly Pro Gly Phe Thr Gly Gly Asp Ile Leu 515 520 525 Arg Arg Thr Asn Thr Gly Gly Phe Gly Pro Ile Arg Val Thr Val Asn 535 Gly Pro Leu Thr Gln Arg Tyr Arg Ile Gly Phe Arg Tyr Ala Ser Thr 545 550 555 Val Asp Phe Asp Phe Phe Val Ser Arg Gly Gly Thr Thr Val Asn Asn 565 570 575 Phe Arg Phe Leu Arg Thr Met Asn Ser Gly Asp Glu Leu Lys Tyr Gly 580 585 Asn Phe Val Arg Arg Ala Phe Thr Thr Pro Phe Thr Phe Thr Gln Ile 595 600 Gln Asp Ile Ile Arg Thr Ser Ile Gln Gly Leu Ser Gly Asn Gly Glu 610 615 620 Val Tyr Ile Asp Lys Ile Glu Ile Ile Pro Val Thr Ala Thr Phe Glu 630 635 Ala Glu Tyr Asp Leu Glu Arg Ala Gln Glu Ala Val Asn Ala Leu Phe 645 650 655 Thr Asn Thr Asn Pro Arg Arg Leu Lys Thr Asp Val Thr Asp Tyr His 660 665 670 Ile Asp Gln Val Ser Asn Leu Val Ala Cys Leu Ser Asp Glu Phe Cys 675 680 685 Leu Asp Glu Lys Arg Glu Leu Leu Glu Lys Val Lys Tyr Ala Lys Arg 690 695 700 Leu Ser Asp Glu Arg Asn Leu Leu Gln Asp Pro Asn Phe Thr Ser Ile 705 710 715 720 Asn Lys Gln Pro Asp Phe Ile Ser Thr Asn Glu Gln Ser Asn Phe Thr

. .

Ser Ile His Glu Gln Ser Glu His Gly Trp Trp Gly Ser Glu Asn Ile
740 745 750

Thr Ile Gln Glu Gly Asn Asp Val Phe Lys Glu Asn Tyr Val Thr Leu 755 760 765

Pro Gly Thr Phe Asn Glu Cys Tyr Pro Thr Tyr Leu Tyr Gln Lys Ile 770 780

Gly Glu Ser Glu Leu Lys Ala Tyr Thr Arg Tyr Gln Leu Arg Gly Tyr 785 790 795 800

Ile Glu Asp Ser Gln Asp Leu Glu Ile Tyr Leu Ile Arg Tyr Asn Ala 805 810 815

Lys His Glu Thr Leu Asp Val Pro Gly Thr Glu Ser Leu Trp Pro Leu 820 825 830

Ser Val Glu Ser Pro Ile Gly Arg Cys Gly Glu Pro Asn Arg Cys Ala 835 840 845

Pro His Phe Glu Trp Asn Pro Asp Leu Asp Cys Ser Cys Arg Asp Gly 850 855 860

Glu Lys Cys Ala His His Ser His His Phe Ser Leu Asp Ile Asp Val 865 870 875 880

Gly Cys Thr Asp Leu His Glu Asn Leu Gly Val Trp Val Val Phe Lys 885 890 895

Ile Lys Thr Gln Glu Gly His Ala Arg Leu Gly Asn Leu Glu Phe Ile
900 905 910

Glu Glu Lys Pro Leu Leu Gly Glu Ala Leu Ser Arg Val Lys Arg Ala 915 920 925

Glu Lys Lys Trp Arg Asp Lys Arg Glu Lys Leu Gln Leu Glu Thr Lys 930 935 940

Arg Val Tyr Thr Glu Ala Lys Glu Ala Val Asp Ala Leu Phe Val Asp 945 950 955 960

Ser Gln Tyr Asp Arg Leu Gln Ala Asp Thr Asn Ile Gly Met Ile His 965 970 975

Ala Ala Asp Lys Leu Val His Arg Ile Arg Glu Ala Tyr Leu Ser Glu 980 985 990

Leu Pro Val Ile Pro Gly Val Asn Ala Glu Ile Phe Glu Glu Leu Glu
995 1000 1005

Gly His Ile Ile Thr Ala Ile Ser Leu Tyr Asp Ala Arg Asn Val 1010 1015 1020

Val Lys Asn Gly Asp Phe Asn Asn Gly Leu Thr Cys Trp Asn Val 1025 1030 1035

Lys Gly His Val Asp Val Gln Gln Ser His His Arg Ser Asp Leu 1040 1045 1050

Val Ile Pro Glu Trp Glu Ala Glu Val Ser Gln Ala Val Arg Val 1055 1060 1065

Cys Pro Gly Cys Gly Tyr Ile Leu Arg Val Thr Ala Tyr Lys Glu 1070 1075 1080

Gly Tyr Gly Glu Gly Cys Val Thr Ile His Glu Ile Glu Asn Asn 1085 1090 1095

Thr Asp Glu Leu Lys Phe Lys Asn Arg Glu Glu Glu Glu Val Tyr 1100 1105 1110

Pro Thr Asp Thr Gly Thr Cys Asn Asp Tyr Thr Ala His Gln Gly 1115 1120 1125

Thr Ala Gly Cys Ala Asp Ala Cys Asn Ser Arg Asn Ala Gly Tyr 1130 1135 1140 Glu Asp Ala Tyr Glu Val Asp Thr Thr Ala Ser Val Asn Tyr Lys 1145 1150 1155

Pro Thr Tyr Glu Glu Glu Thr Tyr Thr Asp Val Arg Asp Asn 1160 1165 1170

His Cys Glu Tyr Asp Arg Gly Tyr Val Asn Tyr Pro Pro Val Pro 1175 1180 1185

Ala Gly Tyr Val Thr Lys Glu Leu Glu Tyr Phe Pro Glu Thr Asp 1190 1195 1200

Thr Val Trp Ile Glu Ile Gly Glu Thr Glu Gly Lys Phe Ile Val 1205 1210 1215

Asp Ser Val Glu Leu Leu Leu Met Glu Glu 1220 1225